

FIG. 1

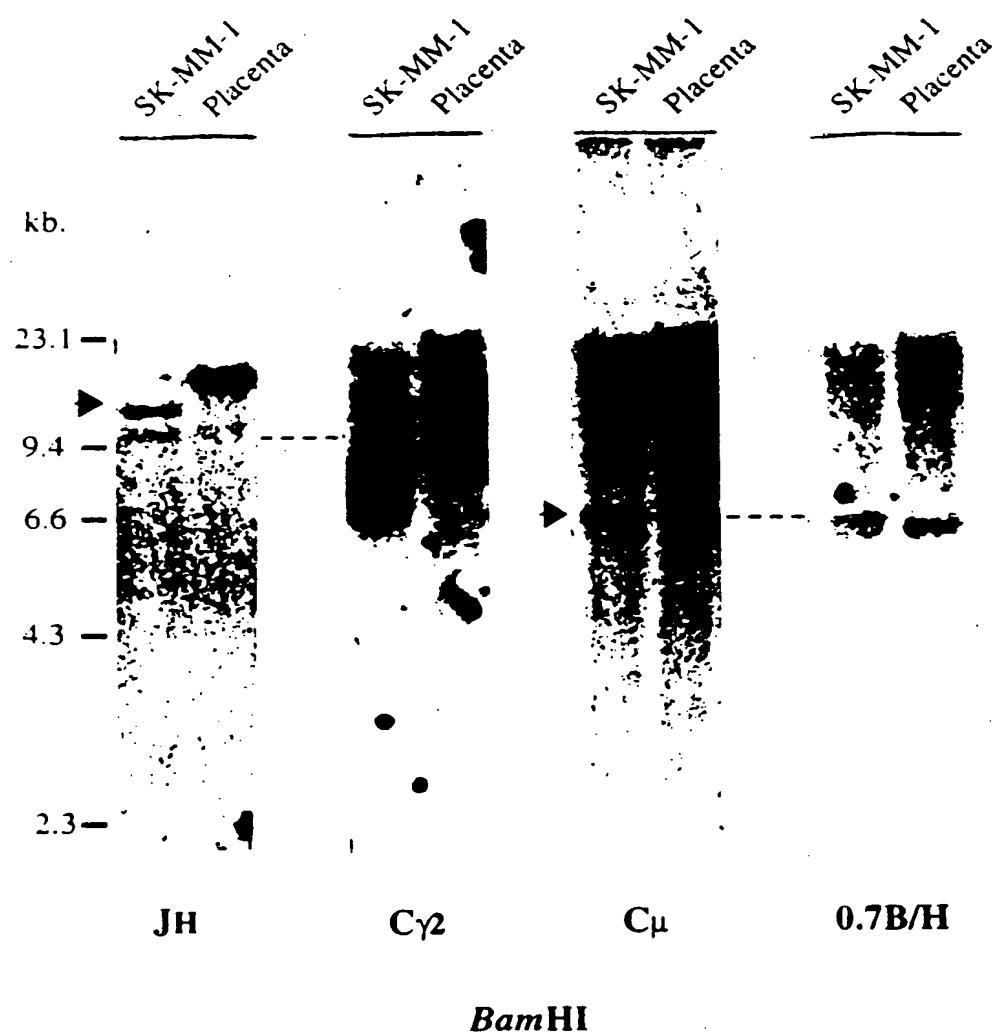
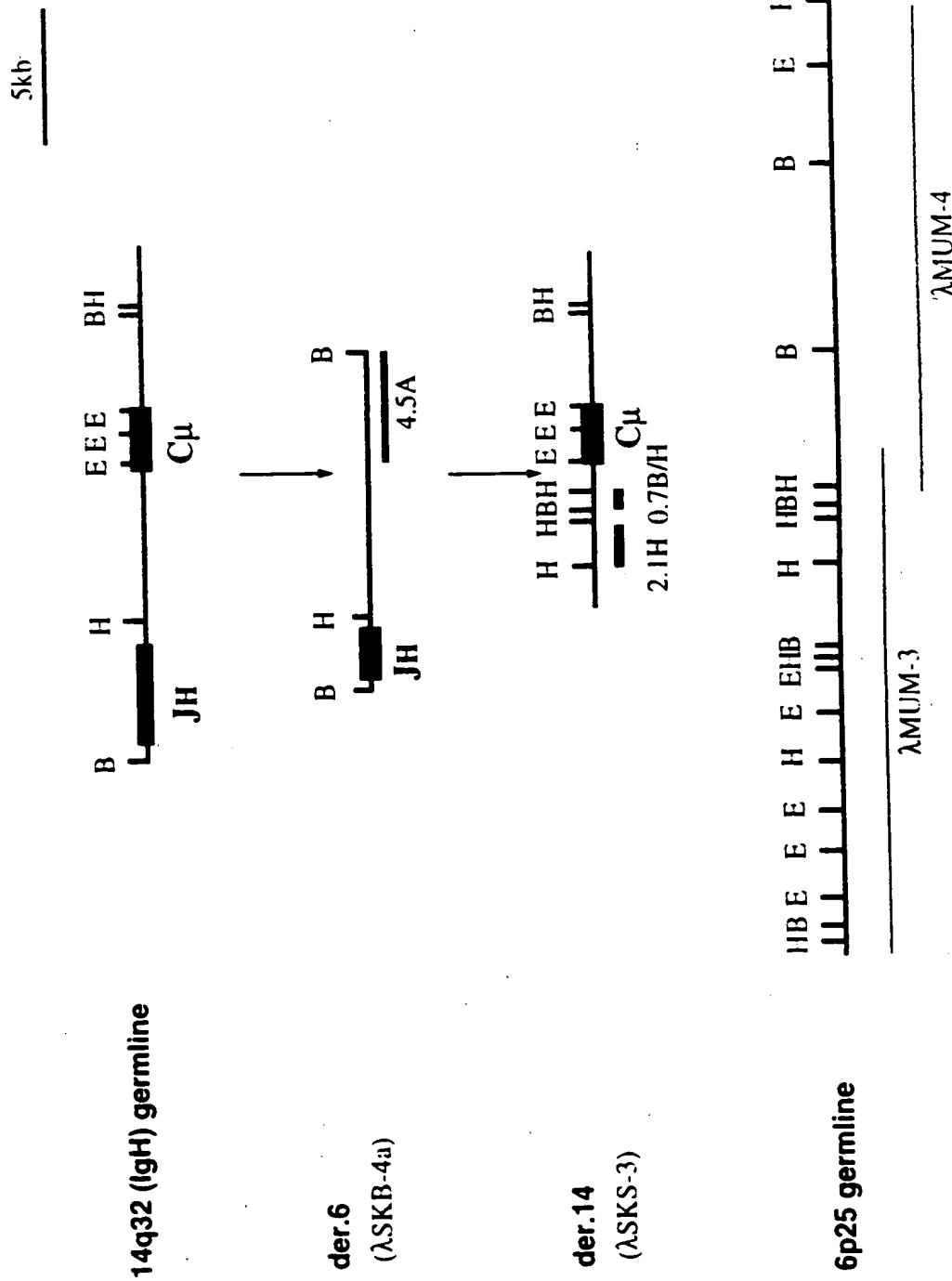


FIG. 2A



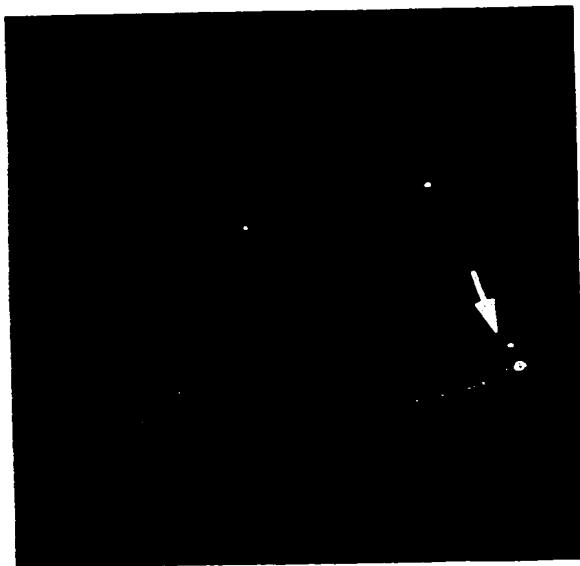
**FIG. 2B**

.6	TTTCTCTACAGTCACCTCCCTGTTACAAAGATAATTACATAAGTCCAGTTACTTACAAACAAAGTTAGT
.6	GGGCTGGCC-TGGTGGGGCAGCCACAGGGGACGGC-AGTAGTGAAGTCCAGTTACTTACAAACAAAGTTAGT
14	GGGCTGGCCTTGGTGGGGCAGCCACAGGGGACGGCAAGTAGTGGGGCACTCAGAACGGCCACTCAGGCCGACAG
	4400 4410 4420 4430 4440 4450 4460 4470

.6	TATTA <del>G</del> GGAA <del>A</del> CTT <del>C</del> AGGATT <del>C</del> AGTCCAGATA <del>A</del> TTTTAAA <del>A</del> ACTCTAAA <del>A</del> ATGGACAGGGCTAGAAT
.14	TATTA <del>G</del> GGAA <del>A</del> CTT <del>C</del> AGGATT <del>C</del> AGGAGGCAT <del>G</del> AGGAGGCAGCTCCCTCACCCCTCCCTCCTTGTAC
.14	GGCACTCAGAACGCCAC <del>T</del> CAGCCCCGACAGGCAGGGCACAGGAGGCAGCTCCCTCACCCCTCCCTCCTTGT--
	4450 4460 4470 4480 4490 4500 4510

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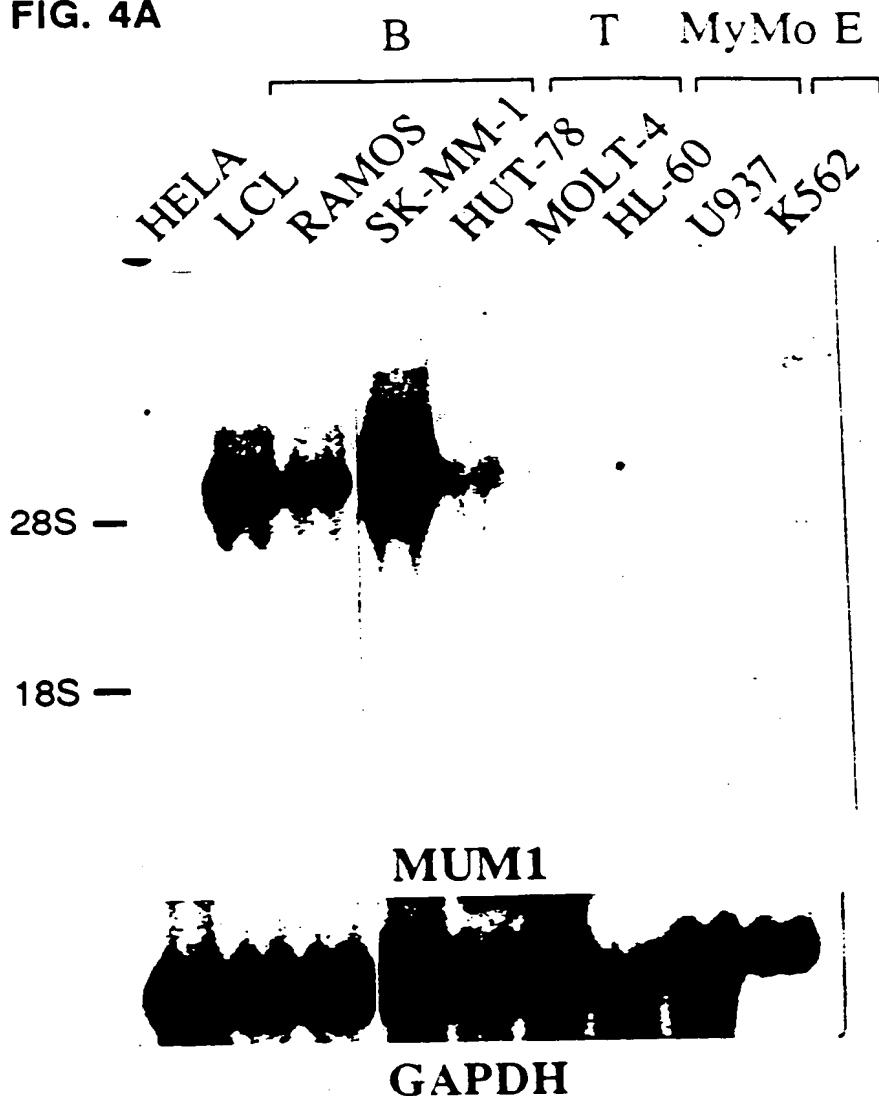
**FIG. 3A**



**FIG. 3B**



**$\lambda$ MUM-3**

**FIG. 4A**

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**FIG. 4B**

PreB    Mature B    Plasma

697    LCL    RAMOS    BJA-B    RPMI-8226  
U-266

28S —

18S —

MUM1

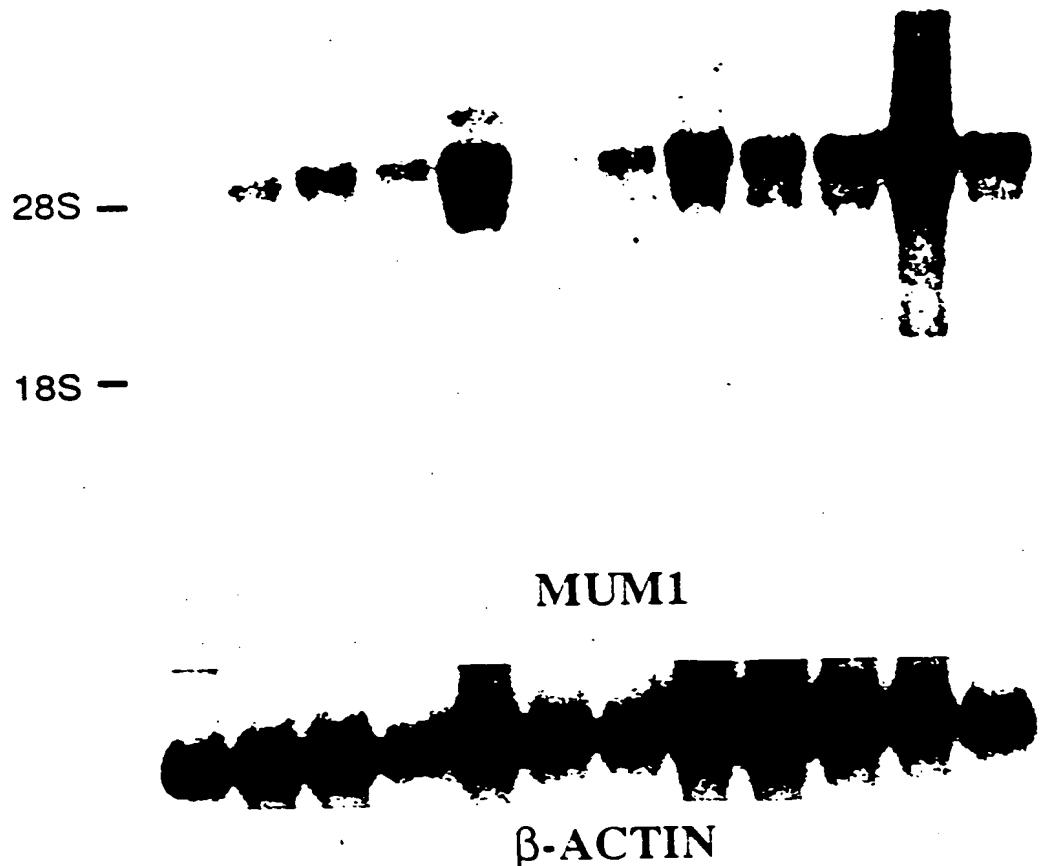
GAPDH

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**FIG. 4C**

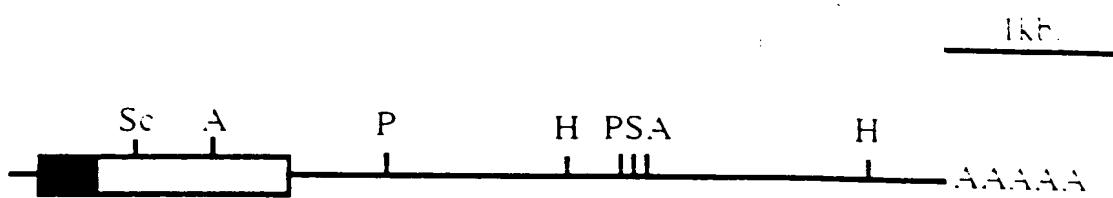
RAMOS  
RPMI-8226  
U-266  
EJM  
SK-MM-1  
XG-1  
XG-2  
XG-4  
XG-5  
XG-6  
XG-7  
XG-10

IL-6 dependent



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**FIG. 5A**



**FIG. 5B-1**

1 GCCTGACCAA CATGGTAAAA CCCCATCTCT GCTAAAAACTA CAAAAAAAT  
51 GCTGGATGTG GTGGCAGGGGA ACCTGTCATC CCAGCTAGTT GGGAGACT  
101 GGCAGGAGAA TCGCTCGATC TTGGGACCCA CCGCTGCCCT CAGCTCCG  
151 TCCAGGGCGA GTGCAGAGCA CAGCGGGCGG AGGACCCCGG GCGCGGGC  
201 GGACGGCACC CGGGGCATGA ACCTGGAGGG CGGCAGGCCGA GGCAGGAGA  
M N L E G G G R G G E  
251 TCGGCATGAG CGCGGTGAGC TGCGGCAACG GGAAGCTCCG CCAGTGGC  
G M S A V S C G N G K L R Q W L  
301 ATCGACCAGA TCGACAGCGG CAAGTACCCC GGGCTGGTGT GGGAGAAC  
I D Q I D S G K Y P G L V W E N  
351 GGAGAAGAGC ATCTTCCGCA TCCCCTGGAA GCACGCGGGC AAGCAGGA  
E K S I F R I P W K H A G K Q D  
401 ACAACCGCGA GGAGGACGCC GCGCTCTTCA AGGCTTGGGC ACTGTTA  
N R E E D A A L F K A W A L F K  
451 GGAAAGTTCC GAGAAGGCAT CGACAAGCCG GACCCCTCCCA CCTGGAAG  
G K F R E G I D K P D P P T W K  
501 GCGCCTGCAG TGCGCTTTGA ACAAGAGCAA TGACTTTGAG GAACTGGT  
R L R C A L N K S N D F E E L V  
551 AGCGGAGCCA GCTGGACATC TCAGACCCGT ACAAAAGTGT A CAGGATTG  
R S Q L D I S D P Y K V Y R I V  
601 CCTGAGGGAG CCAAAAAAGG AGCCAAGCAG CTCACCCCTGG AGGACCCG  
P E G A K K G A K Q L T L E D P  
651 GATGTCCATG AGCCACCCCT ACACCATGAC AACGCCTTAC CCTTCGCT  
M S M S H P Y T M T T P Y P S L  
701 CAGCCCAGCA GGTTCACAAAC TACATGATGC CACCCCTCGA CCGAAGCT  
A Q Q V H N Y M M P P L D R S W  
751 AGGGACTACG TCCCGGATCA GCCACACCCG GAAATCCCGT ACCAATGT  
R D Y V P D Q P H P E I P Y Q C  
801 CATGACGTT GGACCCCGCG GCCACCACTG GCAAGGCCA GCTTGTGA  
M T F G P R G H H W Q G P A C E  
851 ATGGTTGCCA GGTGACAGGA ACCTTTATG CTTGTGCCCG ACCTGAGT  
G C Q V T G T F Y A C A P P E S  
901 CAGGCTCCCG GAGTCCCCAC AGAGCCAAGC ATAAGGTCTG CCGAAGCC  
Q A P G V P T E P S I R S A E A  
951 GGCAGTCTCA GACTGCCGGC TGCACATCTG CCTGTACTAC CGGGAAAT  
A F S D C R L H I C L Y Y R E I  
1001 TCGTGAAGGA GCTGACCAACG TCCAGCCCG AGGGCTGCCG GATCTCCC  
V K E L T T S S P E G C R I S F  
1051 GGACATACGT ATGACGCCAG CAACCTGGAC CAGGTCCGT TCCCCCTAC  
G H T Y D A S N L D Q V L F P Y  
1101 AGAGGACAAT GGCCACAGGA AAAACATTGA GAACCTGCTG AGCCACCI  
E D N G H R K N I E N L L S H L

## FIG. 5B-2

1151 AGAGGGCCGT GGTCCCTCTGG ATGGCCCCCG ACGGGCTCTA TGCAGAAAC  
       R G V V L W M A P D G L Y A K R  
 1201 CTGTGCCAGA GCACGATCTA CTGGGACGGG CCCCTGGCGC TGTGCAAC  
       L C Q S T I Y W D G P L A L C N I  
 1251 CCGGCCAAC AACTGGAGA GAGACCAGAC CTGCAAGCTC TTTGACAC  
       R P N K L E R D Q T C K L F D T  
 1301 AGCAGTTCTT GTCAGAGCTG CAAGCGTTG CTCACCAACGG CCGCTCCC  
       Q F L S E L Q A F A H H G R S L  
 1351 CCAAGATTCC AGGTGACTCT ATGCTTGGA GAGGAGTTTC CAGACCTC  
       P R F Q V T L C F G E E F P D P  
 1401 GAGGCCAAGA AAGCTCATCA CAGCTCACGT AGAACCTCTG CTAGCCAG  
       R Q R K L I T A H V E P L L A R  
 1451 AACTATATTA TTTGCTCAA CAAAACAGTG GACATTCCT GAGGGGCT  
       L Y Y F A Q Q N S G H F I R S Y  
 1501 GATTACCAAG AACACATCAG CAATCCAGAA GATTACCAACA GATCTATC  
       D L P E H I S N P E D Y H R S I  
 1551 CCATTCCTCT ATTCAAGAAT GAAATATGTC AAGATGAGTG GTTTCTT  
       H S S I Q E \*  
 1601 TCCTTTTTTT TTTTTTTTT TTTGATAACGG AGATACGGGG TCTTGCTC  
 1651 TCTCCCAGGC TGGAGTGCAG TGACACAAATC TCAGCTCACT GTGACCTC  
 1701 CCTCCTGGGT TCAAGAGACT CTCCTGCCTC AGCCTCCCTG GTAGCTGG  
 1751 TTACAGGTGT GAGCCACTGC ACCCACCCAA GACAAGTGAT TTTCATTG  
 1801 AATATTGAC TTTAGTGAAA GCGTCCAATT GACTGCCCTC TTACTGTT  
 1851 GAGGAACCTCA GAAGTGGAGA TTTCAGTTCA GCGGTTGAGG AGAATTGC  
 1901 CGAGACAAGC ATGGAAAATC AGTGACATCT GATTGGCAGA TGAGCTTA  
 1951 TCAAAAGGAA GGGTGGCTT GCATTTCTT GTGTTCTGTA GACTGCCA  
 2001 ATTGATGATC ACTGTGAAAA TTGACCAAGT GATGTGTTA CATTACT  
 2051 AATGCGCTCT TTAATTGTT GTAGATTAGG TCTTGCTGGA AGACAGAG  
 2101 AACTGCCTT TCAGTATTGA CACTGACTAG AGTGATGACT GCTTGTAG  
 2151 ATGTCGTGC CATTCTCAG GGAAGTAAGA TGTAAATTGA AGAACGCCT  
 2201 CACGTAAAG AAATGTATTA ATGTATGTAG GAGCTGCAGT TCTTGTGG  
 2251 GACACTTGCT GAGTGAAGGA AATGAATCTT TGACTGAAGC CGTGCCTG  
 2301 GCCTGGGGGA GGCCCATCCC CCACCTGCCA GCGGTTCCCT GGTGTGGG  
 2351 CCTCTGCCCT ACCCTCCTTC CCATTGGCTT TCTCTCCTTG GCCTTCC  
 2401 GAAGCCAGTT AGTAAACTTC CTATTTCTT GAGTCAAAAA ACATGAGC  
 2451 TACTCTGGGA TGGGACATT TTGTCTGTCC TACAATCTAG TAATGTCT  
 2501 GTAATGGTTA AGTTTCTTG TTTCTGCATC TTTTGACCC TCATTCTT  
 2551 GAGATGCTAA AATTCTTCGC ATAAAGAAGA AGAAATTAAG GAACATAA  
 2601 CTTAATACTT GAACTGTTGC CCTTCTGTCC AAGTACTTAA CTATCTGT  
 2651 CCTTCCTCTG TGCCACGCTC CTCTGTTGT TTGGCTGTCC AGCGATCA  
 2701 CATGGCGACA CTAAAGGAGG AGGAGCCGGG GACTCCCAGG CTGGAGAG  
 2751 CTGCCAGGAC CCACCACTGG AAGCAGGATG GAGCTGACTA CGGAACCTG  
 2801 CACTCAGTGG GCTGTTCTG CTTATTTCAT CTGTTCTATG CTTCCTCG  
 2851 CCAATTATAG TTTGACAGGG CCTTAAAATT ACTTGGCTTT TTCCAAAT  
 2901 TTCTATTTAT AGAAATCCCA AAGACCTCCA CTTGCTTAAG TATACCTA  
 2951 ACTTACATTG TTGTGGTTT GAGAAAGTAC AGCAGTAGAC TGGGGCGT

## FIG. 5B-3

3051 TCAGCAGAAG ATTGCGTTAG CTCTCAATG TGTGTTCTG CTTTTCTA  
 3101 GGATATTTA AATTCAATTCA ACAAGCACCT AGTAAGTGCC TGCTGTAT  
 3151 CTACATTACA CAGTCAGCC TTTATCAAGC TTGAGTGAGCA GTGAGCAC  
 3201 AACATTATT TTTTAATGTT TAAAGAGTTT CTAATATTAAG AGTCAGAA  
 3251 TTAATACAAT TAATATTAAT ATTAACATACA GAAAAGACAA ACAGTAGA  
 3301 ACAGCAAAAA AATAAAAAGG ATCTCCTTT TTCCCAGCCC AAATTCTC  
 3351 CTCTAAAAGT GTCCACAAAGA AGGGGTGTTT ATTCTTCCAA CACATTTC  
 3401 TTTTCTGTAA ATATACATAA ACTTAAAAG AAAACCTCAT GGAGTCAT  
 3451 TGCACACACT TTTCATGCAG TGCTCTTGT AGCTAAACAG TGAAGATT  
 3501 CCTCGTTCTG CTCAGAGGCC TTGCTGTGGA GCTCCACTGC CATGTACC  
 3551 GTAGGGTTTG ACATTCATT AGCCATGCAA CATGGATATG TATTGGGC  
 3601 CAGACTGTGT TTGTTGAACT GCAGTGATGT ATACATCTA TAGATGCA  
 3651 GTATTTGGG GTATATTATC CTAAGGGAAAG ATAAAGATGA TATTAAGA  
 3701 TGCTGTTCA CGGGGCCCTT ACCTGTGACC CTCTTGCTG AAGAATAT  
 3751 AACCCCCACAC AGCACTTCAC AGAAGCTGTC TTGGAAGTCT GTCTCAGG  
 3801 CACCCGTCT TCTTAATTCT CCAAGCGGAT GCTCCATTTC AATTGCTT  
 3851 TGACTTCTTC TTCTTTGTTT TTTAAATAT TATGCTGCTT TAACAGTG  
 3901 GCTGAATTTC CTGGAAAATG CTTCTTGGCT GGGGCCACTA CCTCCTTT  
 3951 TATCTTACA TCTATGTGTA TGTGACTTT TTAAAATTCT GAGTGATC  
 4001 GGGTATGACC TAGGGAATGA ACTAGCTATG GAAATAACTC AGGGTTAG  
 4051 ATCCTAGCAC TTGTCTCAGG ACTCTGAAAA GGAACGGCTT CCTCATTG  
 4101 TGTCTTGATA AAGTGGAAATT GGCAAACTAG AATTTAGTTT GTACTCAG  
 4151 GACAGTGCTG TTGAAGATTG GAGGACTTGT TAAAGAGCAC TGGTCAT  
 4201 GGAAAAAAATG TATGTGTCTC CCCAGGTGCA TTTTCTTGGT TTATGTCT  
 4251 TTCTTGAGAT TTTGTATATT TAGGAAACACC TCAAGCAGTA ATTAATATC  
 4301 CCTGGAACAC TATAGAGAAC CAAGTGACCG ACTCATTTC AACTGAAAC  
 4351 TAGGAAGCCC CTGAGTCCTG AGCGAAAACA GGAGAGTTAG TCGCCCTAC  
 4401 GAAAACCCAG CTAGACTATT GGCTATGAAAC TAAAAAGAGA CTGTGCCAC  
 4451 GTGAGAAAAAA TGTAAAATCC TACAGTGGAA TGAGCAGCCC TTACAGTGT  
 4501 GTTACCAACCA AGGGCAGGTA GGTATTAGTG TTTGAAAAAG CTGGTCTT  
 4551 AGCGAGGGCA TAAATACAGC TAGCCCCAGG GGTGGAACAA CTGTGGGAC  
 4601 CTTGGGTACT CGCACCTCTT GGCTTTGTTG ATGCTCCGCC AGGAAGGCC  
 4651 CTTGTGTGTG CGTGTCAAGTT ACTTTTTAG TAACAATTCA GATCCAGTC  
 4701 AAACTTCCGT TCATTGCTCT CCAGTCACAT GCCCCCACCT CCCCACAGG  
 4751 GAAAGTTTTT CTGAAGTGTG GGGATTGGTT AAGGTCTTTA TTTGTATTAA  
 4801 GTATCTCCCC AAGTCCTCTG TGGCCAGCTG CATCTGTCTG AATGGTGCC  
 4851 GAAGGCTCTC AGACCTTACA CACCAATTG TAAGTTATGT TTTACATGC  
 4901 CCGTTTTGA GACTGATCTC GATGCAGGTG GATCTCCTTG AGATCCTGA  
 4951 AGCCTGTTAC AGGAATGAAG TAAAGGTCAAG TTTTTTTGT ATTGATTTT  
 5001 ACAGCTTGA GGAACATGCA TAAGAAATGT AGCTGAAGTA GAGGGGACG  
 5051 GAGAGAAGGG CCAGGCCGGC AGGCCAACCC TCCTCCAATG GAAATTCCCC  
 5101 TGTTGCTTCA AACTGAGACA GATGGGACTT AACAGGCAAT GGGGTCCAC  
 5151 TCCCCCTCTT CAGCATCCCC CGTACC

FIG. 6A

MOM-1 (23-72)	KLROWLIDQI	DSGKYPGLWW	ENEEKSIEFRI	PWKHAGKQDY	NREEDDAALFK
LSIRF (23-72)	KLROWLIDQI	DSGKYPGLWW	ENEEKSIEFRI	PWKHAGKQDY	NREEDDAALFK
IRF-1 (7-56)	RYRPLWEEQI	ENNOIPGLWW	ENEEKMIECI	PWKHAGKQDY	DINKDACLFR
IRF-2 (7-56)	RYRPLWEEQI	ENNTIPGLWW	ENKEKKEFQI	PWKHAGKQDY	DVEKDAPLFR
ICSBP (9-60)	RLRQWLIEQI	DSSMYPGLWW	ENEEKSIEFRI	PWKHAGKQDY	NOEVADASIFK
ISGF3Y (11-60)	YLRNWVVEQY	ESGQEPGVQW	DDTAKTMFRI	PWKHAGKQDF	REDDDAAFFK
IRF-3 (7-55)	RILPMLVSQI	DIGQLEGVWW	VNKSRTFRI	PWKHAGKQDF	AQQEDFG1EQ
MOM-1 (73-122)	AWAIEFKGKFR	EGIDKPDPP	WKTRLRICALN	KSNDFEELVE	RSOLDISDPY
LSIRF (73-122)	AWAIEFKGKFR	EGIDKPDPP	WKTRLRICALN	KSNDFEELVE	RSOLDISDPY
IRF-1 (57-106)	SWAIIHTGRYK	ACEKEPDKT	WKANFRCAVN	SIPDTEEVKD	CRNKGEASAV
IRF-2 (57-106)	NWAIIHTGKHO	PSVDKPDPT	WKANFRCAVN	SLPEEEVKW	KSIIKKGNNAF
ICSBP (59-107)	AWAIEFKGKFR	EC.DKAEPAT	WKTRLRICALN	KSPDFFEVID	RSOLDISDPY
ISGF3Y (61-109)	AWAIEFKGKFR	EG.DTGCQAV	WKTRLRICALN	KSSEEKEVPE	RGMDVAEPY
IRF-3 (56-104)	AWAIEATGAYV	PGRDKPDILPT	WKRNFRSALN	RKEGLRLAED	RSK.DPHDPH
MOM-1 (123-130)	KVYRIVPE				
LSIRF (123-130)	KVYRIVPE				
IRF-1 (107-114)	RVYRMLPP				
IRF-2 (107-114)	RVYRMLPL				
ICSBP (108-115)	KVYRIVPE				
ISGF3Y (110-117)	KVYOLIPP				
IRF-3 (105-112)	KTYEFVNS				

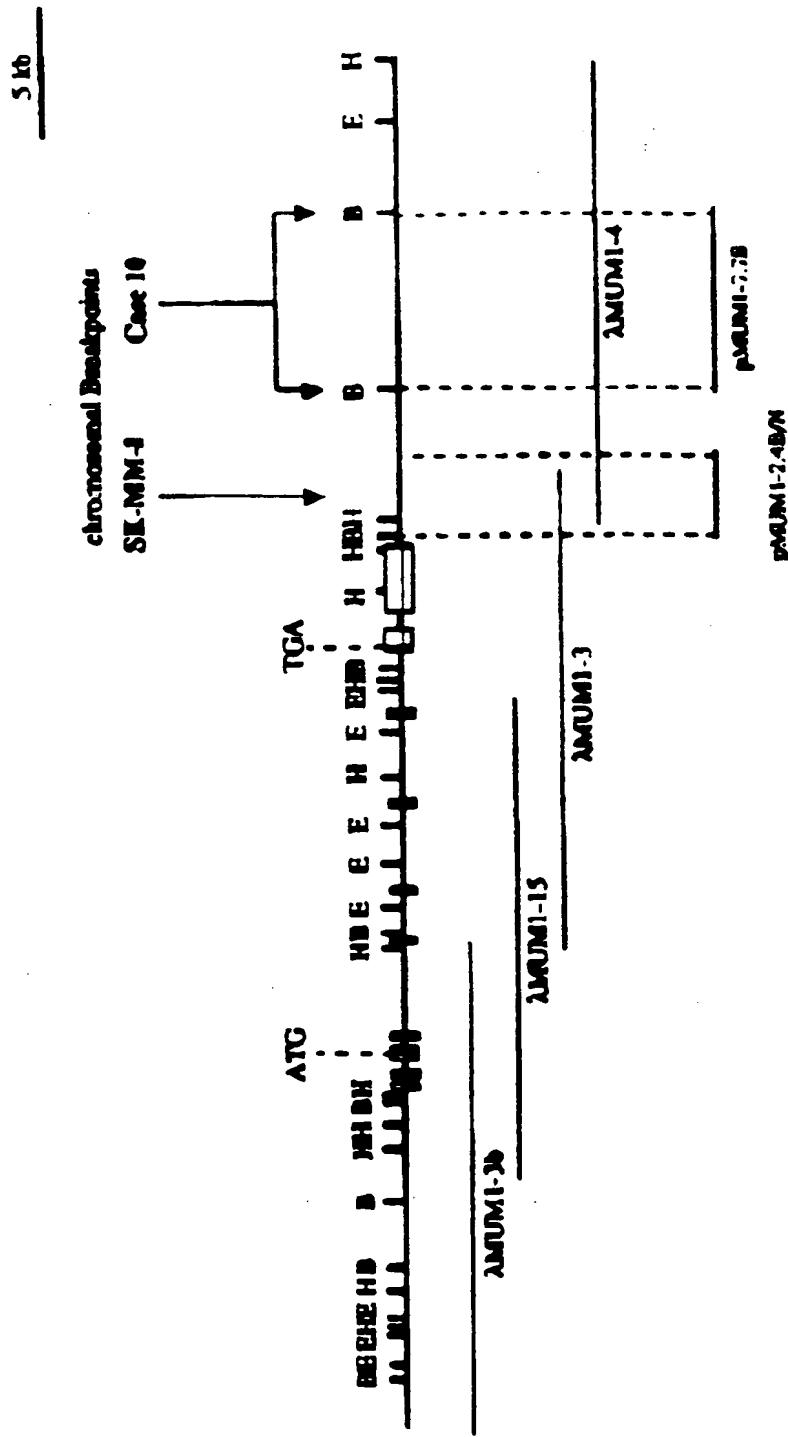
FIG. 6B

MUM-1 (327-372)	KRLCQSTIY	DGPLAL	... ... ...	CNDREVKIER	DOTCKLDTQ	QFLSEQVA
LSIRF (327-372)	KRLCQSRIV	DGPLAL	... ... ...	CSDREVKIER	DOTCKLDTQ	QFLSEQVA
ICSBP (289-334)	KRLCQGRVFC	SGNAVV	... ... ...	CKGRPKIER	DEVQVQTS	OFFRQOQY
ISGF3Y (290-335)	QRLCPPISTK	NAPQAP	... ... ...	PGPGCHLPS	NECEVYERA	YCRDVRVF
IRF-3 (264-333)	QRLGHCHTYK	AVSEELLPNS	... ... ...	KGPDGEVPK	KEGGVILIG	PIVDILIT

MOM-1 (373-421)	HHGRSLPQQ	VTCCEGEEFP	DPQROR.KEL	TAHVSPILLAR	QIYFFAQONS
LSIRF (373-421)	HHGRPAFQQ	WTLCFGEEFP	DPQROR.KEL	TAHVSPILLAR	QIYFFAQONT
ICSBP (335-384)	NSQGRUPDGR	WTCCEGEEFP	DMAPLRSKEL	LVOIQHOLYVR	QIAEAGKSC
ISGF3Y (336-385)	OGLCPPPKFO	WTLCFGEEFP	QSSHTPQNE	TVKNUOAFAK	YILOQTPEQQ
IRF-3 (334-383)	EGSGRSPPRYA	LMTVGEESWP	QDQPWTKRDV	MYKVVPTCIE	ALVEVAFRVGG

### Genomic locus spanning *MUM1* gene

FIG. 7

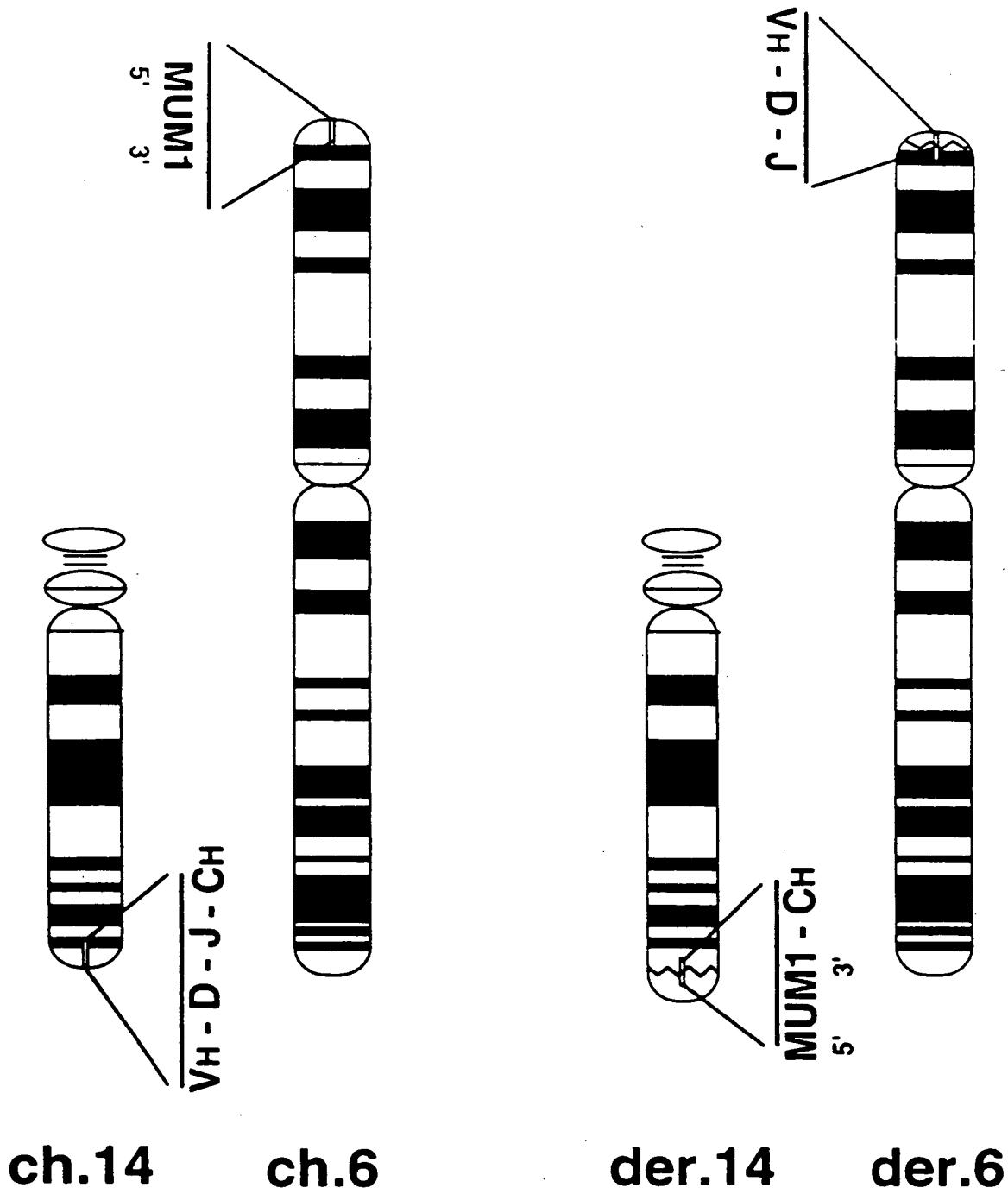


*pMUM1-7.7B* contains the 7.7 kb fragment from the *B/SN* locus.

*B/SN* contains *BamH1*/*NdeI* site is used for cloning.

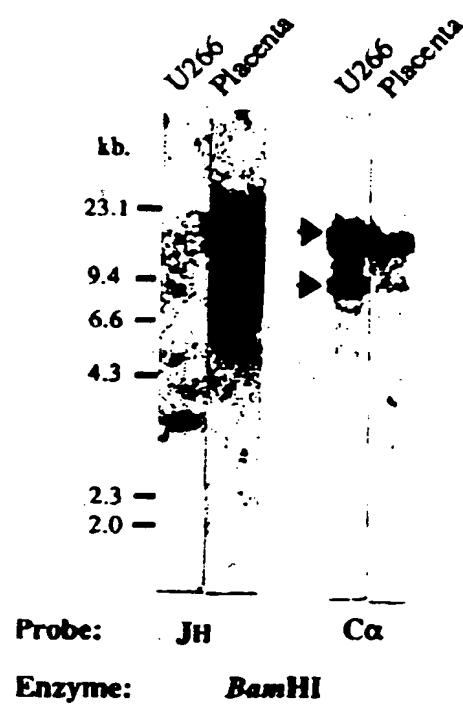
Every genomic insertion is cloned into *pBluescript KS*.

FIG. 8



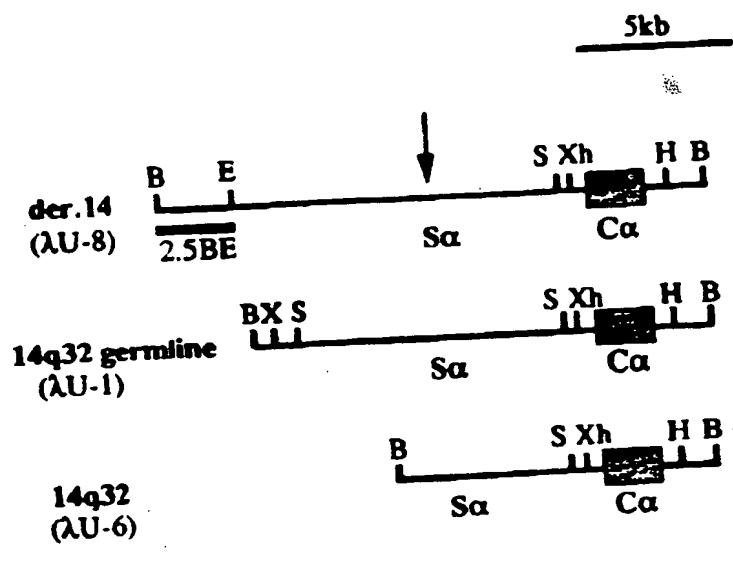
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FIG. 9A



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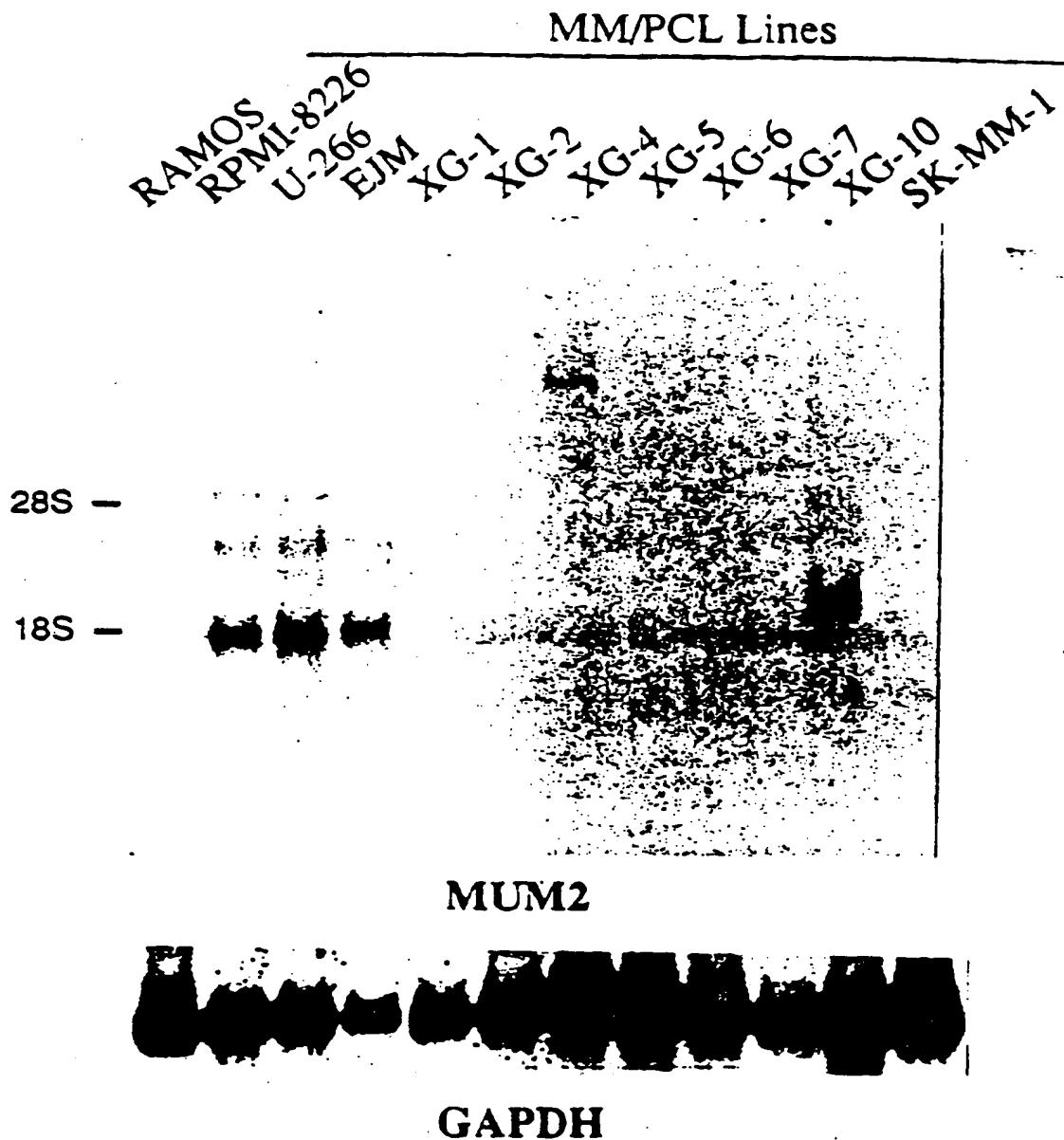
FIG. 9B



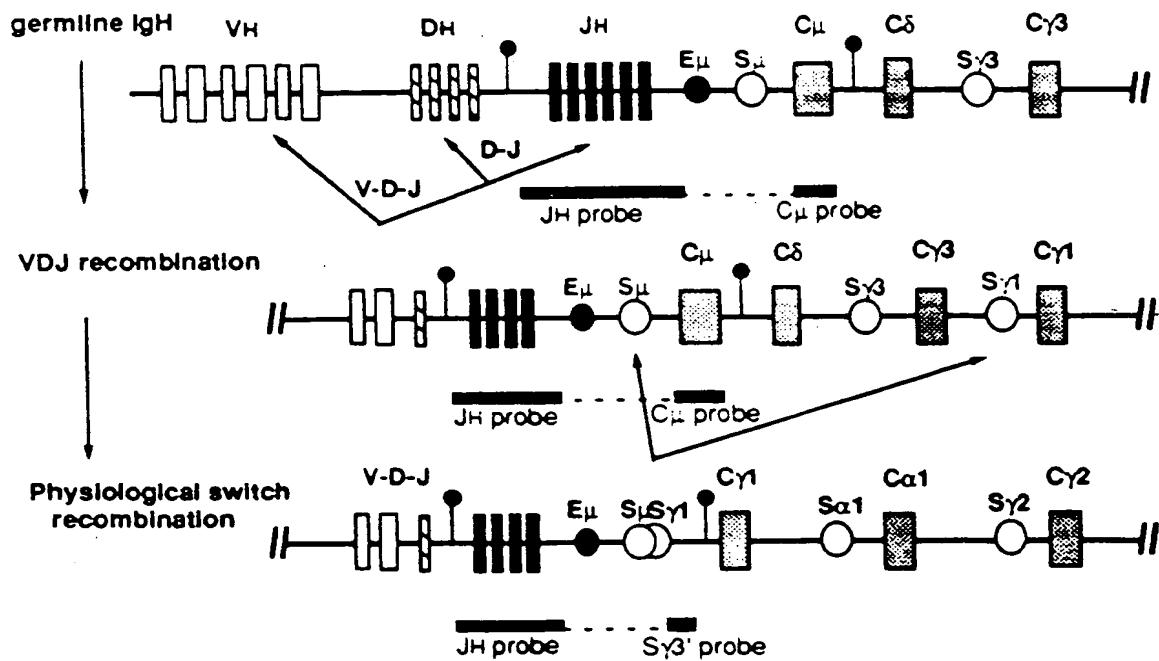
B, *Bam*HI; H, *Hind*III; S, *Sac*II; X, *Xba*I;  
Xh, *Xba*I

↓ chromosomal breakpoint

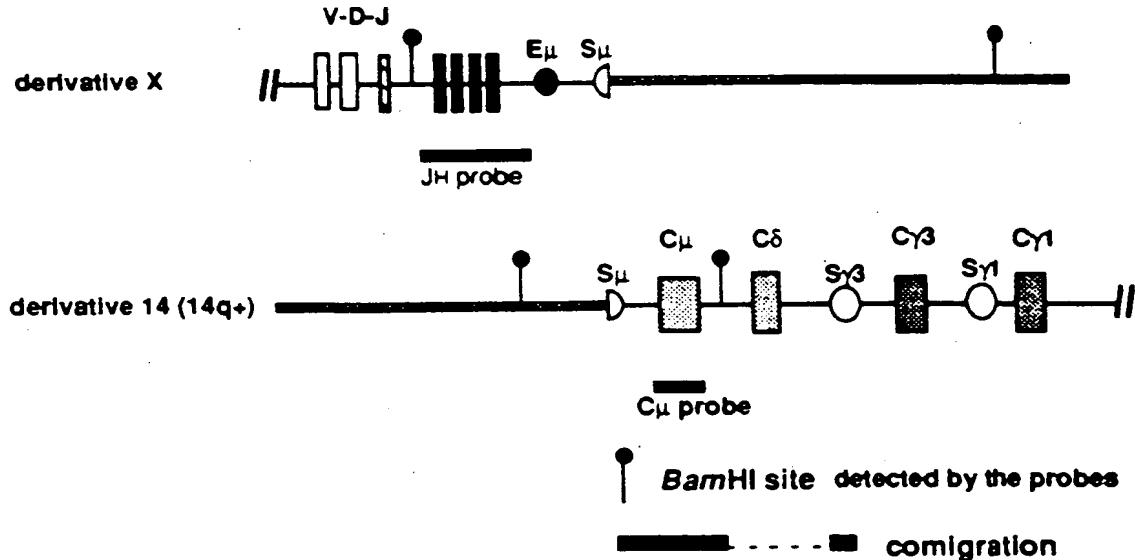
**FIG. 10**  
***MUM2* Transcripts detected in MM/PCL Cell Lines**



**FIG. 11A**  
**Physiological IgH gene rearrangement**

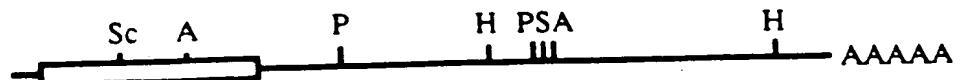


**FIG. 11B**  
**Chromosomal translocation occurring in switch region**



## MUM1 cDNA

1kb.



## pcMUM1-1.6a



Sc; SacII, P;PstI, H;HindIII, S;SacI, A;ApaI

cDNA inserts is cloned into EcoRI / BamHI site of the pBluescript KS+  
 Bacteria strain used is DH5 $\alpha$  cells. pcMUM1-1.6a contains full length open  
 reading frame of nt.217~1572.

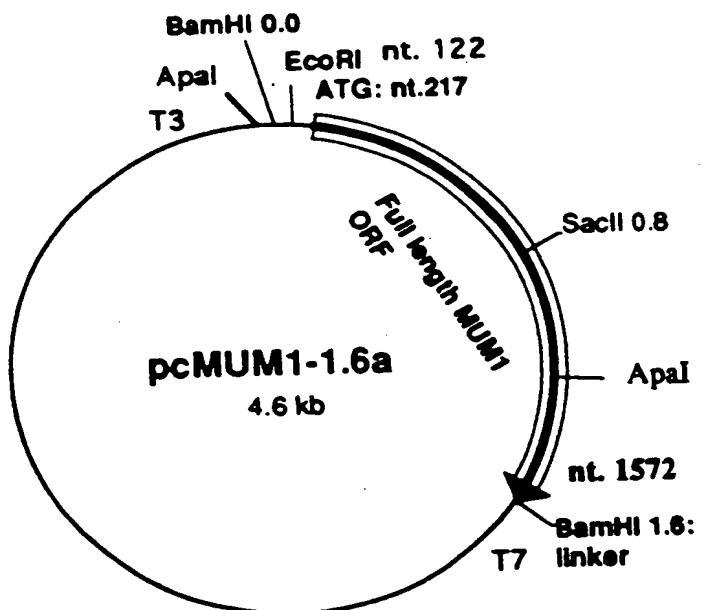
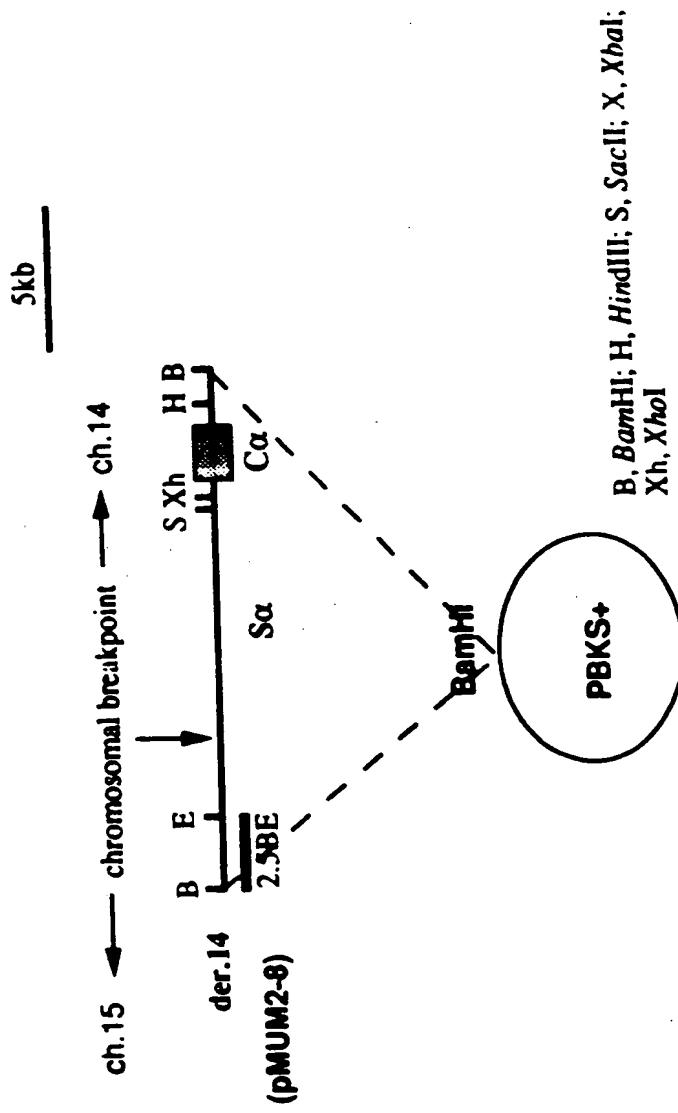


FIG. 12 A-B

## Breakpoint Cloning of the U-266 Cell Line



pMUM2-8 has a 22.0kb insert in BamHI site of pBluescript KS+.

FIG. 13